(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 3 November 2005 (03.11.2005)

(10) International Publication Number WO 2005/103083 A3

(51) International Patent Classification: C07K 16/28 (2006.01) A61P 37/00 (2006.01)

(21) International Application Number:

PCT/IB2005/002476

(22) International Filing Date: 7 February 2005 (07.02.2005)

(25) Filing Language:

English

English

(26) Publication Language:

(30) Priority Data:

60/541,911	6 February 2004 (06.02.2004)	US
60/547,584	26 February 2004 (26.02.2004)	US
60/553,948	18 March 2004 (18.03.2004)	US
60/599,014	6 August 2004 (06.08.2004)	US
60/614,471	1 October 2004 (01.10.2004)	US

(71) Applicant (for all designated States except US): MOR-PHOSYS AG [DE/DE]; Lena-Christ-Str. 48, 82152 Martinsried/Planegg (DE).

(72) Inventors; and

- (75) Inventors/Applicants (for US only): TESAR, Michael [DE/DE]; Karolingerstrasse 26, 82362 Weilheim (DE). JAGER, Ute [DE/DE]; Sammtstrasse 5, 81669 Munchen (DE).
- (74) Agent: JONES DAY, Prinzregentenstrasse 11, 80538 Munich (DE).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): ΛΕ, ΛG, ΛL, ΛΜ,

AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- (88) Date of publication of the international search report: 16 November 2006
- (15) Information about Correction: Previous Correction: see PCT Gazette No. 40/2006 of 5 October 2006

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: ANTI-CD38 HUMAN ANTIBODIES AND USES THEREFOR

(57) Abstract: The present invention provides recombinant antigen-binding regions and antibodies and functional fragments containing such antigen-binding regions that are specific for CD38, which plays an integral role in various disorders or conditions. These antibodies, accordingly, can be used to treat, for example, hematological malignancies such as multiple myeloma. Antibodies of the invention also can be used in the diagnostics field, as well as for investigating the role of CD38 in the progression of disorders associated with malignancies. The invention also provides nucleic acid sequences encoding the foregoing antibodies, vectors containing the same, pharmaceutical compositions and kits with instructions for use. The invention also provides isolated novel epitopes of CD38 and methods of use therefore.



WO 2005/103083 A

INTERNATIONAL SEARCH REPORT

International Application No PCT/IB2005/002476

Relevant to claim No.

A. CLASSIFICATION OF SUBJECT MATTER INV. C07K16/28 A61P37/00

C. DOCUMENTS CONSIDERED TO BE RELEVANT

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) $C\,07K$

Documentation searched other than minimum documentation to the extent that such documents are included. In the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, Sequence Search

Citation of document, with indication, where appropriate, of the relevant passages

	WO 02/06347 A (CRUCELL HOLLAND LOGTENBERG, TON; CILENTI, LUCI/24 January 2002 (2002-01-24) page 2, line 18 - page 3, line claims; examples	1-84		
X	US 2002/164788 A1 (ELLIS JONATHAN HENRY ET AL) 7 November 2002 (2002-11-07) paragraph [0001] - paragraph [0002] paragraph [0069] - paragraph [0072] paragraph [0078] - paragraph [0080] claims; examples -/			
Further documents are listed in the continuation of box C. **Patent family members are listed in annex.** **Patent family members are listed in annex.** **T' later document published after the international filling date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention dilling date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) *O' document referring to an oral disclosure, use, exhibition or other means *P' document published prior to the international filling date but later than the priority date claimed *A' document member of the same patent family Date of the actual completion of the international search *A' document published after the international filling date or priority date and not in conflict with the application but in credit or priority date and not in conflict with the application but in credit or priority date and not in conflict with the application but in credit or priority date and not in conflict with the application but in credit or priority date and not in conflict with the application but credit or priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority d				
Name and n	nailing address of the ISA	Authorized officer		
	European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Rankin, R		

INTERNATIONAL SEARCH REPORT

International Application No PCT/IB2005/002476

		FC1/1B2005/0024/6
C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with Indication, where appropriate, of the relevant passages	Relevant to claim No.
A	ELLIS JONATHAN H ET AL: "Engineered anti-CD38 monoclonal antibodies for immunotherapy of multiple myeloma" JOURNAL OF IMMUNOLOGY, THE WILLIAMS AND WILKINS CO. BALTIMORE, US, vol. 155, no. 2, 1995, pages 925-937, XP002146232 ISSN: 0022-1767 the whole document	1-84
Α	AUSIELLO C M ET AL: "Functional topography of discrete domains of human CD38." TISSUE ANTIGENS. DEC 2000, vol. 56, no. 6, December 2000 (2000-12), pages 539-547, XP002389874 ISSN: 0001-2815 the whole document	1-74
A	HOSHINO S ET AL: "Mapping of the catalytic and epitopic sites of human CD38/NAD+ glycohydrolase to a functional domain in the carboxyl terminus." JOURNAL OF IMMUNOLOGY (BALTIMORE, MD.: 1950) 15 JAN 1997, vol. 158, no. 2, 15 January 1997 (1997-01-15), pages 741-747, XP002389875 ISSN: 0022-1767 abstract	1-84

Form PCT/ISA/210 (continuation of second sheet) (January 2004)

International application No. PCT/IB2005/002476

INTERNATIONAL SEARCH REPORT

Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Although claims 69-74 are directed to a method of treatment of the human body, the search has been carried out and based on the alleged effects of the composition.
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful international Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No
PCT/IB2005/002476

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 0206347	A	24-01-2002	AU EP US	8267501 A 1174440 A1 2003211553 A1	30-01-2002 23-01-2002 13-11-2003
US 2002164788	A1	07-11-2002	NONE		

Form PCT/ISA/210 (patent family annex) (January 2004)